valid OMB control number.

(Modified) PTO/SB/21 (6-98) use through 09/30/2000. OMB 0651-0031

Patent and Trademark fice: U.S. DEPARTMENT OF COMMERCE

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it displays a

TRANSMITTAL FORM

(to be used for all correspondence after initial filing)

Total Number of Pages in This Submission

09/721,507 **Application Number** November 22, 2000 **Filing Date** Joseph A. Affholter **First Named Inventor** Unassigned Group Art Unit Unassigned **Examiner Name** 02-104244US Attorney Docket Number

| | | | ENCLOS | JRES (check all tha | at apply |) | |
|--------------------------------------|----------------------------|---|----------------------------------|--|----------------|-----|---|
| Fee Transmitta | al Form | | Assignme | ent Papers oplication) | | | After Allowance Communication to Group |
| Fee Atta | ached | | Drawing(| s) | | | Appeal Communication to Board of Appeals and Interferences |
| Amendment / | Response | | Licensing | -related Papers | | | Appeal Communication to Group (Appeal Notice, Brief, Reply Brief) |
| After F | Final | | Petition Rand Acco | outing Slip (PTO/SB/6) companying Petition | ⁵⁹⁾ | | Proprietary Information |
| Affidav | vits/declaration(s) | | | o Convert to a al Application | | | Status Letter |
| Extension of 1 | Time Request | | Power of Change of Address | Attorney, Revocation of Correspondence | | X | Additional Enclosure(s) (please identify below): |
| Express Aban | ndonment Request | | | Disclaimer tity Statement | | | receipt acknowledgment postcard |
| X Information Di | isclosure Statement | | | for Refund | | | |
| Certified Copy Document(s) | - | Authorization to Charge Deposit Account Please charge Deposit Account No. 50-0893 for any additional fees associated with this paper or during the pendency of this application, including any extensions of time | | | | | |
| Incomplete A | Missing Parts/ Application | for cor | nsideration | of the documents encl | osed. | | |
| Respo | onse to Missing | Re | emarks | | | | |
| | under 37 CFR | | | | | | |
| | SIGNATU | IRE O | F APPLIC | ANT, ATTORNEY, | OR A | GEI | NT |
| Firm <i>or</i> Individual name | | | | | | | s of Jonathan Alan Quine |
| Signature | Uph Ann | fert | / | | | | |
| Date | June 1, 2001 | | | | | | |

| CERTIFICATE OF MAILING | | | | | | | |
|----------------------------|---|--------------------|----------------------------------|--|--|--|--|
| I hereby certify that this | correspondence is being deposited with the Unite Assistant Commissioner for Patents, Washington | ed States Postal S | ervice as first class mail in an | | | | |
| envelope addressed to: | Assistant Commissioner for Patertis, Washington | 1, 0.0. 20201 0111 | Julie 1. 2001 | | | | |
| Typed or printed name | Alexandra Allison | | | | | | |
| Signature | alexandra alli | Date | June 1, 2001 | | | | |





I hereby certify that this correspondence is being deposited with the United States Postal Service as first class mail in an envelope addressed to:

Assistant Commissioner for Patents, Washington, D.C. 20231, On June 1, 2001

The Law Offices of Jonathan Alan Quine

Alexandra Allison

The Law Offices of Johannah Alah Quine

RECEIVED

JUN 0 7 2001

TECH CENTER 1600/2900

Attorney Docket No. 02-104244US

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re application of:

Joseph A. Affholter, et al.

Application No.: 09/721,507

Filed: November 22, 2000

For: Single-Stranded Nucleic acid
Template-Mediated recombination and

nucleic acid fragment isolation

Examiner: Unassigned

Art Unit: Unassigned

SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT UNDER 37

CFR § 1.97 and § 1.98

Assistant Commissioner for Patents Washington , D.C. 20231

Sir:

The references cited on attached PTO-1449 forms are being called to the attention of the Examiner to make of record references cited in parent application USSN 09/656,549 filed September 6, 2000. Pursuant to 37 CFR § 1.98(d), copies of references cited in parent application USSN 09/656,549 filed September 6, 2000 are not provided. However the applicants will gladly provided fresh copies of any references requested by the Examiner. It is respectfully requested that the cited information on the previously submitted 1449 forms

Application No.: 09/721,507

Page 2

for application USSN 09/656,549 filed September 6, 2000 be expressly considered during the prosecution of this application, and that references be made of record therein and appear among the "references cited" on any patent to issue therefrom.

As provided for by 37 CFR 1.97(g) and (h), no inference should be made that the information and references cited are prior art merely because they are in this statement and no representation is being made that a search has been conducted or that this statement encompasses all the possible relevant information.

Applicant believes that <u>no fee is required</u> for submission of this statement, since it is being submitted prior to the first Office Action. However, if a fee is required, the Commissioner is authorized to deduct such fee from the undersigned's Deposit Account No. 50-0893. Please deduct any additional fees from, or credit any overpayment to, the abovenoted Deposit Account.

Respectfully submitted,

Christopher C. Sappenfield

Reg. No. 45,073

The Law Offices of Jonathan Alan Quine

P.O. Box 458

Alameda, CA 94501

Tel: (510) 337-7871

Fax: (510) 337-7877

CCS:afa

(Modified) PTO/SB/08A-B (10-96) Approved for use through 10/31/99. OMB 0651-0031

stitute for form 1449A-B/PTO

INFORMATION DISCLOSURE STATEMENT BY APPLICANT

| | | | | S. PATENT DOCUMENTS | Date of Publication of | Pages, Columns, lines, | |
|----------------------|-------------|--------------------------|-------------------------|--|---------------------------|---|-------|
| Examiner Initials | Cite No. | U.S. Patent Do Number | Kind Code (if known) | Name of Patentee or Applicant of Cited Document | Cited Document MM-DD-YYYY | Where Relevant Passages or Relevant Figures Appeal | |
| | AA | 5,756,316 | | Schellenberger | 05-26-1998 | | |
| | AB | 5,783,431 | | Peterson et al. | 07-21-1998 | RECEN | VE |
| | AC | 5,798,208 | | Crea | 08-25-1998 | 11111.6 | |
| | AD | 5,824,485 | | Thomspon, et al. | 10-20-1998 | | 001 |
| | AE | 5,830,650 | | Crea | 11-03-1998 | TECH CENTER 16 | 00/29 |
| | AF | 6,057,103 | | Short | 05-02-2000 | | |
| | AG | 6,165,793 | | Stemmer | 12-26-2000 | | |
| | АН | 6,168,919 | | Short | 01-02-2001 | | |
| | Al | 6,171,820 | | Short | 01-09-2001 | | |
| | AJ | 6,174,673 | | Short, et al. | 01-16-2001 | | |
| | AK | 6,180,406 | | Stemmer | 1-30-2001 | | |

| | | | | FOREIGN | PATENMT DOCUMEN | ITS | | |
|----------------------|-------------|--------|----------------------|--------------------------------|--|--|---|---|
| Examiner Initials | Cite No. | Office | Foreign Patent Docum | ent Kind Code (if known) | Name of Patentee or Applicant of Cited Document | Date of Publication of Cited Document MM-DD-YYYY | Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear | т |
| | AL | EP | 0527809 | | Crea, Roberto | 08-19-1995 | | |
| | AM | wo | 95/22625 | | Affymax Tech. N.V. | 08-24-1995 | | |
| | AN | wo | 96/33207 | | Glaxo Group Limited | 10-24-1996 | | |
| | AO | wo | 97/33957 | | Amway Corp. | 09-18-1997 | | |
| | AP | wo | 98/42727 | | SRI International | 10-01-1998 | | |
| | AQ | wo | 99/29902 | | California Institute of Technology | 06-17-1999 | | |
| | AR | wo | 00/42560 | | Maxygen, Inc. | 07-20-2000 | | |
| | AS | wo | 00/42561 | | Maxygen, Inc. | 07-20-2000 | | |
| | AT | wo | 00/46344 | | Diversa Corp. | 08-10-2000 | | |
| | AU | wo | 00/53744 | | Diversa Corp. | 09-14-2000 | | |
| | AV | wo | 00/58517 | | Diversa Corp. | 10-05-2000 | | |

| | | |
|-----------|------------|--|
| Examiner | Date | |
| Signature | Considered | |

^{*}EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

Sussitute for form 1449A-B/PTO

INFORMATION DISCLOSURE STATEMENT BY APPLICANT

| | | OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS TECH CENT | FR 1 | 600/29 | |
|--------------------------|-------------|---|------|--------|--|
| Examin er Initials | Cite No. | Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published. | Т | 000,23 | |
| | AW | Arnold (1993) "Protein engineering for unusual environments" Current Opinion in Biotechnology 4:450-455 | | | |
| | AX | Arkin & Youvan (1992) Proc. Natl. Acad. Sci. USA 89:7811-7815 | | | |
| | AY | Arkin and Youvan (1992) "Optimizing nucleotide mixtures to encode specific subsets of amino acids for semi-random mutagenesis" <i>Biotechnology</i> 10:297-300 | | | |
| | AZ | Bass et al. (1988) "Mutant Trp repressors with new DNA-binding specificities" Science 242:240-245) | | | |
| | ВА | Botstein & Shortle (1985) "Strategies and applications of in vitro mutagenesis" Science 229:1193-1201 | | | |
| | ВВ | Breyer and Sauer (1989) "Mutational analysis of the fine specificity of binding of monoclonal antibody 51F to lambda repressor" <i>J. Biol. Chem.</i> 264:13355-60 | | | |
| | вс | Caldwell et al. (1992) PCR Methods Applic. 2:28-33 | ļ | | |
| | BD | Carter (1986) "Site-directed mutagenesis" Biochem. J. 237:1-7 | | | |
| | BE | Carter (1987) "Improved oligonucleotide-directed mutagenesis using M13 vectors" <i>Methods in Enzymol.</i> 154: 382-403 | | | |
| | BF | Carter et al. (1985) "Improved oligonucleotide site-directed mutagenesis using M13 vectors" Nucl. Acids Res. 13: 4431-4443 | | | |
| | BG | Dale et al. (1996) "Oligonucleotide-directed random mutagenesis using the phosphorothioate method" <i>Methods Mol. Biol.</i> 57:369-374 | | | |
| | вн | delCardayre, S.B. and Raines, R.T. (1995) Anal. Biochem. 225, 176 | | | |
| | ВІ | Delegrave and Youvan (1993) Biotechnology Research 11:1548-1552 | | | |
| | BJ | Eghtedarzadeh & Henikoff (1986) "Use of oligonucleotides to generate large deletions" <i>Nucl. Acids Res.</i> 14: 5115 | | | |
| | ВК | Fritz et al. (1988) "Oligonucleotide-directed construction of mutations: a gapped duplex DNA procedure without enzymatic reactions in vitro" <i>Nucl. Acids Res.</i> 16: 6987-6999 | | | |
| | BL | Grundström et al. (1985) "Oligonucleotide-directed mutagenesis by microscale 'shot-gun' gene synthesis" <i>Nucl. Acids Res.</i> 13: 3305-3316 | | | |
| | ВМ | Guzman et al. (1995) "Tight regulation, modulation, and high-level expression by vectors containing the arabinose PBAD promoter" <i>J. Bacteriol.</i> 177(14):4121-4130 | | | |
| | BN | Jirholt et al. (1998) "Exploiting sequence space: shuffling in vivo formed complementarity determining regions into a master framework" <i>Gene</i> 215: 471 | | | |
| | | | | _ | |

| Examiner | Date | |
|-----------|------------|--|
| Signature | Considered | |

^{*}EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

Superitute for form 1449A-B/PTO

INFORMATION DISCLOSURE STATEMENT BY APPLICANT

| Complete if Known | | | | | |
|------------------------|---------------------------------|--|--|--|--|
| Application Number | 09/656549 09/721,507 | | | | |
| Filing Date | September 6, 2000 \ \-22-00 | | | | |
| First Named Inventor | Joseph A. Affholter | | | | |
| Group Art Unit | 1645 | | | | |
| Examiner Name | Unassigned | | | | |
| Attorney Docket Number | 02-104240US | | | | |

| E | Johnson, M.G. (1996) Epicentre Forum 3(4),7 |
|-----|--|
| E | Kramer & Fritz (1987) <i>Methods in Enzymol.</i> "Oligonucleotide-directed construction of mutations via gapped duplex DNA" 154:350-367 |
| E | Q Kramer et al. (1984) "Point Mismatch Repair" Cell 38:879-887 |
| E | R Kramer et al. (1984) "The gapped duplex DNA approach to oligonucleotide-directed mutation construction" <i>Nucl. Acids Res.</i> 12: 9441-9456 |
| E | S Kramer et al. (1988) "Improved enzymatic in vitro reactions in the gapped duplex DNA approach to oligonucleotide-directed construction of mutations" <i>Nucl. Acids Res.</i> 16: 7207 |
| E | T Kunkel (1985) "Rapid and efficient site-specific mutagenesis without phenotypic selection" Proc. Natl. Acad. Sci. USA 82:488-492 |
| E | U Kunkel (1987) "The efficiency of oligonucleotide directed mutagenesis" in <i>Nucleic Acids & Molecular Biology</i> (Eckstein, F. and Lilley, D.M.J. eds., Springer Verlag, Berlin) |
| E | V Kunkel et al. (1987) "Rapid and efficient site-specific mutagenesis without phenotypic selection" <i>Methods in Enzymol.</i> 154, 367-382 |
| Е | W Leung et al. (1989) <i>Technique</i> 1:11-15 |
| E | X Lim and Sauer (1991) "The role of internal packing interactions in determining the structure and stability of a protein" <i>J. Mol. Biol.</i> 219:359-76 |
| Е | Y Ling et al. (1997) "Approaches to DNA mutagenesis Anal Biochem. 254(2): 157-178 |
| E | Z Mandecki, Oligonucleotide-directed double-strand break repair in plasmids of <i>Escherichia coli</i> : a method for site-specific mutagenesis" <i>Proc. Natl. Acad. Sci. USA</i> , 83:7177-7181 |
| C | A Meador, J and Kennell, D. (1990) Gene 95:1 |
| C | B Meador, J. et al. (1990) <i>Eur J. Biochem.</i> 187:549 |
| C | Nakamaye & Eckstein (1986) "Inhibition of restriction endonuclease Nci I cleavage by phosphorothicate groups and its application to oligonucleotide-directed mutagenesis" <i>Nucl. Acids Res.</i> 14: 9679-9698 |
| C | Nambiar et al. (1984) "Total synthesis and cloning of a gene coding for the ribonuclease S protein" Science 223:1299-1301 |
| C | Nikiforov et al. (1994) "The Use of Phosphorotioate Primers and Exonuclease Hydrolysis for the Preparation of Single-Stranded PCR Products and their Detection by Solid-Phase Hybridization," PCR Methods and Applications 3:285-291 |
| C | Ostermeier et al. (1999) "A combinatorial approach to hybrid enzymes independent of DNA homology" Nature Biotech 17:1205 |
| . (| Ostermeier et al. (1999) "Combinatorial Protein Engineering by Incremental Truncation," <i>Proc. Natl. Acad. Sci. USA</i> , 96: 3562-67 |
| | |

| | The state of the s | | |
|-----------|--|------------|--|
| Examiner | | Date | |
| Signature | | Considered | |

^{*}EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

OLD E SCORE

Substitute for form 1449A-B/PTO

INFORMATION DISCLOSURE STATEMENT BY APPLICANT

Complete if Known

Application Number

Filing Date

First Named Inventor

Group Art Unit

Examiner Name

Attorney Docket Number

Complete if Known

69/656549

99/72/387

September 6, 2000

1-22-00

1645

Unassigned

2-104240US

| | СН | Ostermeier et al. (1999), "Incremental Truncation as a Strategy in the Engineering of Novel Biocatalysts," <i>Biological and Medicinal Chemistry</i> , 7: 2139-44 |
|---|----|--|
| | CI | Porter et al. (1997) Nucleic Acids Res. 25(8):1611-1617 |
| | CJ | Reidhaar-Olson et al. (1988) Science, 241:53-57 |
| | СК | Reidhaar-Olson et al. (1991) Methods Enzymol. 208:564-86 |
| | CL | Sakamar and Khorana (1988) "Total synthesis and expression of a gene for the a-subunit of bovine rod outer segment guanine" Nucl. Acids Res. 14: 6361-6372 |
| | СМ | Sayers et al. (1988) "Strand specific cleavage of phosphorothioate-containing DNA by reaction with restriction endonucleases" Nucl. Acids Res. 16: 803-814 |
| | CN | Smith (1985) "In vitro mutagenesis" Ann. Rev. Genet. 19:423-462 |
| | со | Stemmer et al., (1995) "Single-step assembly of a gene and entire plasmid form large numbers of oligodeoxy-ribonucleotides" <i>Gene</i> , 164:49-53 |
| | СР | Taylor et al. (1985) "The rapid generation of oligonucleotide-directed mutations at high frequency using phosphorothioate-modified DNA" <i>Nucl. Acids Res.</i> 13: 8765-8787 (1985) |
| | CQ | Taylor et al. (1985) "The use of phosphorothioate-modified DNA in restriction enzyme reactions to prepare nicked DNA" <i>Nucl. Acids Res.</i> 13: 8749-8764 |
| | CR | Wells et al. (1985) "Cassette mutagenesis: an efficient method for generation of multiple mutations at defined sites" <i>Gene</i> 34:315-323 |
| | CS | Wells et al. (1986) "Importance of hydrogen-bond formation in stabilizing the transition state of subtilisin" <i>Phil. Trans. R. Soc. Lond.</i> A 317: 415-423 |
| | СТ | Zhao and Arnold (1997) "Functional and nonfunctional mutations distinguished by random recombination of homologous genes," <i>Proc. Natl. Acad. Sci. U.S.A.</i> 94(15):7997-8000 |
| | CU | Zhao, et al., "Molecular evolution by staggered extension process (StEP) in vitro recombination," Nature Biotechnology (March 1998), 16(3):258-61 |
| | CV | Zhou et al., (1995) Biotechniques 19:34-35 |
| | CW | Zhou et al., (1998) "Regulatory Roles of the P Domain of the Subtilisin-like Prohormone Convertases," <i>J. Biol. Chem.</i> , 273(18):11107 |
| | СХ | Zoller & Smith (1982) "Oligonucleotide-directed mutagenesis using M13-derived vectors: an efficient and general procedure for the production of point mutations in any DNA fragment" Nucleic Acids Res. 10:6487-6500 |
| | CY | Zoller & Smith (1983) "Oligonucleotide-directed mutagenesis of DNA fragments cloned into M13 vectors" <i>Methods in Enzymol.</i> 100:468-500 |
| , | CZ | Zoller & Smith (1987) "Oligonucleotide-directed mutagenesis: a simple method using two oligonucleotide primers" <i>Methods in Enzymol.</i> 154:329-350 |

| | | |
|-----------|------------|--|
| Examiner | Date | |
| Signature | Considered | |

^{*}EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.